From the INTERNATIONAL SEARCHING AUTHORITY

To: SARAH A. KAGAN

BANNER & WITCOFF, LTD. 11TH FLOOR 1001 G STREET, NW WASHINGTON, DC 20001-4597	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION (PCT Rule 44.1)			
	Date of Mailing (day/month/year)			
Applicant's or agent's file reference 03482.00010	FOR FURTHER ACTION See paragraphs 1 and 4 below			
International application No. PCT/US03/25614	International filing date (day/month/year) 15 August 2003 (15.08.2003)			
Applicant GENZYME CORPORATION				
	earch report has been established and is transmitted herewith.			
Filing of amendments and statement under Article 19 The applicant is entitled, if he so wishes, to amend the	!			
When? The time limit for filing such amendments international search report.	is normally two months from the date of transmittal of the			
Where? Directly to the International Bureau of WII 1211 Geneva 20. Switzerland. Facsimile N				
For more detailed instructions, see the notes on the	accompanying sheet.			
2. The applicant is hereby notified that no international sea Article 17(2)(a) to that effect is transmitted herewith.	arch report will be established and that the declaration under			
3. With regard to the protest against payment of (an) add	ditional fee(s) under Rule 40.2, the applicant is notified that:			
applicant's request to forward the texts of both the	been transmitted to the International Bureau together with the e protest and the decision thereon to the designated Offices. applicant will be notified as soon as a decision is made.			
4. Reminders				
Shortly after 18 months from the priority date, the internatio applicant wishes to avoid or postpone publication, a notice of	onal application will be published by the International Bureau. If the withdrawal of the international application, or of the priority claim, is. I and 90 bis. 3. respectively, before the completion of the technical			
Within 19 months from the priority date, but only in respect of some designated Offices, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later); otherwise the applicant must, within 20 months from the priority date, perform the prescribed acts for entry into the national phase before those designated Offices.				
In respect of other designated Offices, the time limit of 30 months (or later) will apply even if no demand is filed within 19 months.				
See the Annex to Form PCT/IB/301 and, for details about th <i>Guide</i> , Volume II, National Chapters and the WIPO Internet si	ne applicable time limits. Office by Office, see the PCT Applicant's te.			
Nome and will also call YALVIO	Authorized officers and			
Name and mailing address of the ISA/US Mail Stop PCT, Attn: ISA/ US Commissioner for Patents	Authorized officer. Francisco Christopher J Nichols, Ph.D.			
P.O. Box 1450 Alexandria, Virginia 22313- 1450 Faccipile No. 671, 273, 2301	Telephone No. (571) 272-1600			
Facsimile No. (571) 273-3201 Form PCT/ISA/220 (April 2002)	(See notes on accompanying sheet)			

PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY

PCT To: SARAH A. KAGAN BANNER & WITCOFF, LTD. 11TH FLOOR NOTIFICATION OF TRANSMITTAL OF 1001 G STREET, NW THE INTERNATIONAL SEARCH REPORT WASHINGTON, DC 20001-4597 OR THE DECLARATION (PCT Rule 44.1) 27 DEC 2005 Date of Mailing (day/month/year) Applicant's or agent's file reference FOR FURTHER ACTION See paragraphs 1 and 4 below 03482.00010 International filing date International application No. (day/month/year) PCT/US03/25614 15 August 2003 (15.08.2003) Applicant GENZYME CORPORATION The applicant is hereby notified that the international search report has been established and is transmitted herewith. Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the international application (see Rule 46): The time limit for filing such amendments is normally two months from the date of transmittal of the international search report. Where? Directly to the International Bureau of WIPO. 34, chemin des Colombettes 1211 Geneva 20. Switzerland. Facsimile No.: (41-22) 338.82.70. For more detailed instructions, see the notes on the accompanying sheet. The applicant is hereby notified that no international search report will be established and that the declaration under Article 17(2)(a) to that effect is transmitted herewith. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that: the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices. no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made. Reminders Shortly after 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in Rules 90 bis.1 and 90 bis.3, respectively, before the completion of the technical preparations for international publication. Within 19 months from the priority date, but only in respect of some designated Offices, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later): otherwise the applicant must. within 20 months from the priority date, perform the prescribed acts for entry into the national phase before those designated Offices. In respect of other designated Offices, the time limit of 30 months (or later) will apply even if no demand is filed within 19

Name and mailing address of the ISA/US Mail Stop PCT, Attn: ISA/ US Commissioner for Patents	Christopher J Nichols, Ph.D.
P.O. Box 1450 Alexandria, Virginia 22313-1450 Faccinita No. (571) 272-3201	Telephone No. (571) 272-1600

See the Annex to Form PCT/IB/301 and, for details about the applicable time limits. Office by Office, see the PCT Applicant's

Form PCT/ISA/220 (April 2002)

Guide, Volume II. National Chapters and the WIPO Internet site.

(See notes on accompanying sheet)

PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference 03482.00010	FOR FURTHER ACTION	see Notific Report (Fo item 5 belo				
International application No. PCT/US03/25614	International filing date (day/month/year) 15 August 2003 (15.08.2003)		(Earliest) Priority Date (day/month/year) 15 August 2002 (15.08.2002)			
Applicant GENZYME CORPORATION						
This international search report has been applicant according to Article 18. A continuous applicant according to Article 18.	opy is being transmitted to the Inte	earching /ernational	Authority and is transmitted to the Bureau.			
This international search report consists of a total of sheets. It is also accompanied by a copy of each prior art document cited in this report.						
Basis of the Report a. With regard to the language, the international search was carried out on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.						
Authority (Rule 23.1(b)). b. With regard to any nucleotid			e international application furnished to this te international application, the international			
	nal application in written form.	Julia fam				
1	rnational application in computer rea this Authority in written form.	idable for				
1 ==	this Authority in computer readable	form.				
the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.						
the statement that the information been furnished.	the statement that the information recorded in computer readable form is identical to the written sequence listing has					
2. Certain claims were four	nd unsearchable (See Box I).					
4. With regard to the title,	omitted by the applicant					
	the text is approved as submitted by the applicant. the text has been established by this Authority to read as follows:					
5. With regard to the abstract,						
	the text is approved as submitted by the applicant.					
the text has been establish may, within one month fr	ned, according to Rule 38.2(b), by the om the date of mailing of this intern	nis Authori ational sea	ity as it appears in Box III. The applicant rch report, submit comments to this Authority.			
6. The figure of the drawings to be	published with the abstract is Figure	No				
as suggested by the applic			None of the figures			
because the applicant faile						
because this figure better characterizes the invention.						

Form PCT/ISA/210 (first sheet) (July 1998)

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/25614

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)			
This international rep	ort has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:		
1. Claim because	Nos.: e they relate to subject matter not required to be searched by this Authority, namely:		
2. Claim becaus such a	Nos.: e they relate to parts of the international application that do not comply with the prescribed requirements to n extent that no meaningful international search can be carried out, specifically:		
3. Claim becaus 6.4(a).	Nos.: te they are dependent claims and are not drafted in accordance with the second and third sentences of Rule		
Box II Observati	ions where unity of invention is lacking (Continuation of Item 2 of first sheet)		
This International Se Please See Continua	earching Authority found multiple inventions in this international application, as follows: tion Sheet		
search	required additional search fees were timely paid by the applicant, this international search report covers all nable claims. searchable claims could be searched without effort justifying an additional fee, this Authority did not invite		
paym	ent of any additional fee.		
3. As or cover	lly some of the required additional search fees were timely paid by the applicant, this international search report s only those claims for which fees were paid, specifically claims Nos.:		
restri	equired additional search fees were timely paid by the applicant. Consequently, this international search report is cted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-13 [drawn to delta(transloconiated protein delta)]		
Remark on Protes	The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.		
	INO protest accompanied the payment of additional seaton rees.		

Form PCT/ISA/210 (continuation of first sheet(1)) (July 1998)

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/25614

A. CLASSIFICATION OF SUBJECT MATTER					
IPC(7) : C12Q 1/00, 1/68, 1/70; G01N 33/53, 33/567, 33/569 US CL : 435/4, 5, 6, 7.1, 7.2, 7.21					
	International Patent Classification (IPC) or to both na	tional classification and IPC			
	DS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols) U.S.: 435/4, 5, 6, 7.1, 7.2, 7.21					
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched					
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Please See Continuation Sheet					
C. DOC	UMENTS CONSIDERED TO BE RELEVANT				
Category *	Citation of document, with indication, where ap		Relevant to claim No.		
A	WANG, Z. ET AL. Cloning and molecular characterization of a human ortholog of Monodelphis TRAP in ultraviolet B-induced melanoma. Melonoma Research. April 2004. Vol. 14, No. 2, pp. 107-114, entire document.				
A	MIYAZAKI, K. ET AL. NEDL1, a Novel Ubiquitin-protein Isopeptide Ligase for Dishevelled-1, Targets Mutant Superoxide Dismutase-1. The Journal of Biological Chemistry. 19 March 2004, Vol. 279, No. 12, pages 11327-11335, entire document.				
A	BRENNER, V. ET AL. Genomic Organization of Two Novel Genes on Human Xq28: Compact Head to Head Arrangement of IDHgamma and TRAPdelta is Conserved in Rat and Mouse. Genomics. 15 August 1997, Vol. 44, No. 1, pages 8-14, entire document.				
A	KUNST, C.B. ET AL. Mutations in SOD1 associate cause novel protein interactions. Nature Genetics. Ja 94, entire document.		1-13		
Further	documents are listed in the continuation of Box C	See patent family annex.			
"A" document of particu	pecial categories of cited documents: t defining the general state of the art which is not considered to be ular relevance plication or patent published on or after the international filing	"T" later document published after the integrate and not in conflict with the applicate and not in conflict with the applicate principle or theory underlying the document of particular relevance; the considered novel or cannot be considered."	ication but cited to understand 2 invention 3 claimed invention cannot be ered to involve an inventive		
	t which may throw doubts on priority claim(s) or which is cited to the publication date of another citation or other special reason (as	"Y" document of particular relevance; the considered to involve an inventive ste combined with one or more other such as a particular relevance.	e claimed invention cannot be ep when the document is th documents, such combination		
"O" document	t referring to an oral disclosure, use, exhibition or other means	being obvious to a person skilled in t			
	"P" document published prior to the international filing date but later than the				
Date of the a	Date of the actual completion of the international search Date of mailing of the internation 2 search 2009.				
15 February 2005 (15.02.2005) Name and mailing address of the ISA/US Authorized officer Authorized officer					
Mai Cor	Mail Stop PCT. Attn: ISA/US Commissioner for Patents Christopher J Nichols, Ph.D.				
Ale	D. Box 1450 xandria, Virginia 22313-1450 D. (571) 273-3201	Telephone No. (571) 273-1600			

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INTERNATIONAL SEARCH REPORT

C. (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT				
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
		Relevant to claim No. 1-13		
		ů.		

PCT/US03/25614

INTERNATIONAL SEARCH REPORT

BOX II. OBSERVATIONS WHERE UNITY OF INVENTION IS LACKING

This application contains the following inventions or groups of inventions which are not so linked as to form a single general inventive concept under PCT Rule 13.1. In order for all inventions to be examined, the appropriate additional examination fees must be paid.

- Group 1, claim(s) 1-13, drawn to a method to aid in diagnosing glioma comprising the steps of detecting an expression product.
- Group 2, claim(s) 14-21, drawn to a method of treating a glioma comprising the steps of administering an antibody.
- Group 3, claim(s) 22-35 and 47-60, drawn to a method of identifying a test compound as a potential anti-caner or anti-glioma drug.
- Group 4, claim(s) 36-46, drawn to a method to aid in diagnosing glioma comprising the steps of detecting an mRNA.
- Group 5, claim(s) 61-67, drawn to a method to induce an immune response to glioma.

This application contains claims directed to more than one species of the generic invention. These species are deemed to lack unity of invention because they are not so linked as to form a single general inventive concept under PCT Rule 13.1. The species listed do not share a common structure or function.

The following claim(s) are generic:

- Group 1, Claim 1, all species listed therein.
- Group 2, Claim 14, all species listed therein.
- Group 3, Claims 22 and 47, all species listed therein in each claim.
- Group 4, Claim 36, all species listed therein.
- Group 5, Claim 61, all species listed therein.

In order for more than one species to be examined, the appropriate additional examination fees must be paid.

Applicant is invited to select one species per generic claim listed per group elected.

The inventions listed as Groups 1-5 do not relate to a single general inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons:

- Group 1 is drawn to the special technical feature of detecting an expression product, which is not shared by any of the other groups.
- Group 2 is drawn to the special technical feature of administering an antibody, which is not shared by any of the other groups.
- Group 3 is drawn to the special technical feature of identifying a test compound, which is not shared by any of the other groups.
- Group 4 is drawn to the special technical feature of detecting an mRNA, which is not shared by any of the other groups.
- Group 5 is drawn to the special technical feature of inducing an immune response, which is not shared by any of the other groups.

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Continuation of B. FIELDS SEARCHED Item 3:	
WEST (USPT, PGPUBS, USOCR, JPO, EPO, DERWENT); NCBI (PUBMED);	STN (BIOSCIENCE)
delta(translocon-associated protein delta); colorectal tumor, colon tumor, colon can	ncer, colorectal cancer, gene expression, ESTs,
microarrays	
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